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Applicants:

Mark Ledeboer et al.

Application No.:

10/700,333

UEU 2 1 2006

# AMENDMENTS TO THE CLAIMS

Please replace all prior versions and listings of claims with the amended claims as follows:

1. (Currently amended) A compound of formula I:

I

or a pharmaceutically acceptable salt thereof, wherein:

- R<sup>1</sup> is a phenyl, cyclohexyl, cyclopentyl, pyridyl, morpholino, piperazinyl, or piperidinyl group, wherein R<sup>1</sup> Q /<sub>F</sub><sup>‡</sup>,
- Q is a C<sub>1-2</sub> alkylidene chain wherein one methylene unit of Q is optionally replaced by O, NR, NRCO, NRCONR, NRCO<sub>2</sub>, CO, CO<sub>2</sub>, CONR, OC(O)NR, SO<sub>2</sub>, SO<sub>2</sub>NR, NRSO<sub>2</sub>, NRSO<sub>2</sub>NR, C(O)C(O), or C(O)CH<sub>2</sub>C(O);
- Arth is a 5-7 membered saturated, particulty unsaturated, or fully unsaturated monocyclic ring having 0-3 hetero items independently selected from nitrogen, oxygen, or sulfur, or an 8-12 membered saturated, partially unsaturated, or fully unsaturated bicyclic ring system having 0-5 heteroatoms independently selected from nitrogen, oxygen, or sulfur; v/herein Arth is optionally substituted with q independent occurrences of Z-RZ; wherein q is 0-5, Z is a bond or is a C1-C6 alkylidene chain wherein up to two non-adjacent methylene units of Z are optionally and independently replaced by CO, CO2, COCO, CONR, OCONR, NRNR, NRNRCO, NRCO, NRCO2, NRCONR, SO, SO2, NRSO2, SO2NR, NRSO2NR, O, S, or NR; and each occurrence of RZ is independently selected from R', halogen, NO2, CN, OR', SR', N(R')2, NR'COR', NR'CON(R')2,

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NR'CO<sub>2</sub>R', COR', CO<sub>2</sub>R', OCOR', CON(R')<sub>2</sub>, OCON(R')<sub>2</sub>, SOR', SO<sub>2</sub>R', SO<sub>2</sub>N(R')<sub>2</sub>, NR'SO<sub>2</sub>R', NR'SO<sub>2</sub>N(R')<sub>2</sub>, COCOR', or COCH<sub>2</sub>COR';

each occurrence of R is independently hydrogen or an optionally substituted C<sub>1-6</sub> aliphatic group; and each occurrence of R' is independently hydrogen or an optionally substituted C<sub>1-6</sub> aliphatic group, a 3-8-membered saturated, partially unsaturated, or fully unsaturated monocyclic ring having 0-3 heteroatoms independently selected from nitrogen, oxygen, or sulfur, or an 8-12 membered saturated, partially unsaturated, or fully unsaturated bicyclic ring system having 0-5 heteroatoms independently selected from nitrogen, oxygen, or sulfur; or R and R', two occurrences of R, or two occurrences of R', are taken together with the atom(s) to which they are bound to form an optionally substituted 3-12 membered saturated, partially unsaturated, or fully unsaturated monocyclic or bicyclic ring having 0-4 heteroatoms independently selected from nitrogen, oxygen, or sulfur;

 $Z^{l}$  is N;

 $Z^{7}$  is  $C(U)_{n}R^{Y}$ ;

T and U are each independently a bond or a saturated or unsaturated C<sub>1-6</sub> alkylidene chain, wherein up to two methylene units of the chain are optionally and independently replaced by CO, CC<sub>2</sub>, COCO, CONR, OCONR, NRNR, NRNRCO, NRCO, NRCO<sub>2</sub>, NRCONR, SO, SO<sub>2</sub>, NRSO<sub>2</sub>, SO<sub>2</sub>NR, NRSO<sub>2</sub>NR, O, S, or NR; m and π are each independently 0 or 1;

RX and RY are each independently selected from R or Ar1;

Z<sup>2</sup> is N or CR<sup>2</sup>; Z<sup>3</sup> is N or CR<sup>3</sup>; Z<sup>4</sup> is N or CR<sup>4</sup>; Z<sup>5</sup> is N or CR<sup>5</sup>; and Z<sup>6</sup> is N or CR<sup>6</sup>, wherein each occurrence of R<sup>2</sup>, R<sup>3</sup>, R<sup>4</sup>, R<sup>5</sup> or R<sup>6</sup> is independently R<sup>U</sup> or (V)<sub>p</sub>R<sup>V</sup>, provided that a) no more than three of Z<sup>2</sup>, Z<sup>3</sup>, Z<sup>4</sup>, Z<sup>5</sup> or Z<sup>6</sup> is N, and b) at least one of Z<sup>3</sup>, Z<sup>4</sup> or Z<sup>5</sup> is CR<sup>3</sup>, CR<sup>4</sup>, or CR<sup>5</sup>, respectively, and at least one of R<sup>3</sup>, R<sup>4</sup>, or R<sup>5</sup> is R<sup>U</sup>.

each occurrence of R<sup>U</sup> is NRCOR<sup>7</sup>, CONR(R<sup>7</sup>), SO<sub>2</sub>NR(R<sup>7</sup>), NRSO<sub>2</sub>R<sup>7</sup>,

NRCONR(R<sup>7</sup>), NRSO<sub>2</sub>NR(R<sup>7</sup>), or CONRNR(R<sup>7</sup>), wherein R<sup>7</sup> is (CH<sub>2</sub>)<sub>r</sub>-Y-R<sup>8</sup>, and
t is 0, 1, or 2, Y is a bond or is O, S, NR<sup>9</sup>, -OCH<sub>2</sub>-, -SCH<sub>2</sub>, -NR<sup>9</sup>CH<sub>2</sub>, O(CH<sub>2</sub>)<sub>2</sub>-, -

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S(CH<sub>2</sub>)<sub>2</sub>, or -NR<sup>9</sup>(CH<sub>2</sub>)<sub>2</sub>, and R<sup>8</sup> is Ar<sup>2</sup>, or R<sup>8</sup> and R<sup>9</sup>, taken together with the nitrogen atom, form an optionally substituted 5-8 membered heterocyclyl or heteroaryl ring having 1-3 heteroatoms independently selected from nitrogen, oxygen or sulfur;

each occurrence of V is a bond or a saturated or unsaturated C<sub>1.6</sub> alkylidene chain, wherein up to two methylene units of the chain are optionally and independently replaced by CO, CO<sub>2</sub>, COCO, COINR, OCONR, NRNR, NRNRCO, NRCO, NRCO<sub>2</sub>, NRCONR, SO, SO<sub>2</sub>, NRSO<sub>2</sub>, SO<sub>2</sub>NR, NRSO<sub>2</sub>NR, O, S, or NR;

each occurrence of p is 0 or 1;

each occurrence of RV is R or Ar2; and

Ar<sup>2</sup> is a 5-7 membered saturated, partially unsaturated, or fully unsaturated monocyclic ring having 0-3 heteroatoms independently selected from nitrogen, oxygen, or sulfur, or an 8-12 membered saturated, partially unsaturated, or fully unsaturated bicyclic ring system having 0-5 heteroatoms independently selected from nitrogen, oxygen, or sulfur; wherein Ar<sup>2</sup> is optionally substituted with r independent occurrences of W-R<sup>W</sup>; wherein r is 0-3, W is a bond or is a C<sub>1</sub>-C<sub>6</sub> alkylidene chain wherein up to two non-adjacent methylene units of W are optionally replaced by CO, CO<sub>2</sub>, COCO, CONR, OCONR, NRNR, NRNRCO, NRCO, NRCO<sub>2</sub>, NRCONR, SO, SO<sub>2</sub>, NRSO<sub>2</sub>, SO<sub>2</sub>NR, NRSO<sub>2</sub>NR, O, S, or NR; and each occurrence of R<sup>W</sup> is independently selected from R', halogen, NO<sub>2</sub>, CN, OR', SR', N(R')<sub>2</sub>, NR'COR', NR'CON(R')<sub>2</sub>, NR'CO<sub>2</sub>R', COR', CO<sub>2</sub>R', OCOR', CON(R')<sub>2</sub>, OCON(R')<sub>2</sub>, SOR', SO<sub>2</sub>R', SO<sub>2</sub>N(R')<sub>2</sub>, NR'SO<sub>2</sub>R', NR'SO<sub>2</sub>N(R')<sub>2</sub>, COCOR', or COCH<sub>2</sub>COR';

#### provided that:

a) when Z<sup>7</sup> is CH and ring B is phenyl and at least one of R<sup>3</sup> or R<sup>4</sup> is NHCOR<sup>7</sup>, then R<sup>1</sup> is not phenyl only substituted with two or three occurrences of OR'; and
 b) when Z<sup>7</sup> is CH and ring B is phenyl and at least one of R<sup>3</sup> of R<sup>4</sup> is NHCOR<sup>7</sup>, SO<sub>2</sub>R<sup>7</sup>, CONRR<sup>7</sup>, then R<sup>1</sup> is not phenyl only substituted with one occurrence of CON(R')<sub>2</sub> in the para position.

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#### 2-4. (Canceled)

- 5. (Currently amended) The compound of claim 1, wherein R<sup>1</sup> is an optionally substituted from phenyl, cyclohexyl, or pyridyl group.
- 6. (Original) The compound of claim 1, wherein R<sup>1</sup> is optionally substituted phenyl.
- 7. (Original) The compound of claim 1, wherein q is 0, 1, 2, or 3 and each independent occurrence of ZR<sup>Z</sup> is C<sub>1-4</sub>alkyl, N(R')<sub>2</sub>, OR', SR', CON(R')<sub>2</sub>, NR'COR', NR'SO<sub>2</sub>R', or SO<sub>2</sub>N(R')<sub>2</sub>.
- 8. (Original) The compound of claim 1, wherein q is 1 and ZR<sup>z</sup> is -NH<sub>2</sub>, -OH, C<sub>1</sub>.

  4alkoxy, or -S(O)<sub>2</sub>NH<sub>2</sub>.
- 9. (Original) The compound of claim 1, wherein q is 1, and ZR<sup>Z</sup> is in the meta position and ZR<sup>Z</sup> is -NH<sub>2</sub>, -OH, C<sub>1-4</sub>alkoxy, or -S(O)<sub>2</sub>NH<sub>2</sub>.
- 10. (Original) The compound of claim 1, wherein  $(T)_m R^X$  and  $(U)_n R^Y$  are hydrogen, halogen, NO<sub>2</sub>, CN, OR, SR or N(R)<sub>2</sub>, or C<sub>1-4</sub>aliphatic optionally substituted with oxo, OR, SR, N(R)<sub>2</sub>, halogen, NC<sub>2</sub> or CN.
- 11. (Original) The compound of claim 1, wherein  $(T)_m R^X$  and  $(U)_n R^Y$  are each independently hydrogen, Me, OH, ONle or  $N(R)_2$ .
- 12. (Original) The compound of claim 1, wherein  $(T)_m R^X$  and  $(U)_n R^Y$  are each hydrogen.
- 13. (Original) The compound of claim 1, wherein ring B is one of rings i-xiv:

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14. (Original) The compound of claim 1, wherein t is 0, Y is a bond, and R<sup>8</sup> is an optionally substituted aryl or heteroaryl moiety.

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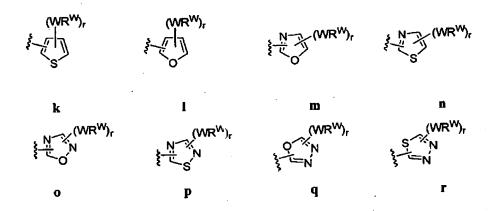
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- 15. (Original) The compound of claim 1, wherein t is 0, Y is a bond, and R<sup>8</sup> is an optionally substituted heteroaryl moiety.
- 16. (Original) The compound of claim 1, wherein R<sup>7</sup> is -CH<sub>2</sub>-Y-R<sup>8</sup>, and Y is NR<sup>9</sup>, O or S, and R<sup>8</sup> is an optionally substituted aryl or heteroaryl moiety.
- 17. (Original) The compound of claim 1, wherein R<sup>7</sup> is -CH<sub>2</sub>-Y-R<sup>8</sup>, and Y is NR<sup>9</sup>, O or S, and R<sup>8</sup> is an optionally substituted aryl moiety.
- 18. (Original) The compound of claim 1, wherein t is 0 or 1, Y is NR<sup>9</sup>, and R<sup>8</sup> and R<sup>9</sup>, taken together with the nitrogen atom, form a 5-8 membered heterocyclyl or heteroaryl ring having 1-3 heteroatoms independently selected from nitrogen, oxygen or sulfur.
- 19. (Original) The compound of claim 1, wherein R<sup>8</sup> is a 5- or 6-membered aryl or heteroaryl group having one of the formulae:

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20. (Original) The compound of claim 1, wherein R<sup>8</sup> is a 5- or 6-membered heteroaryl group having one of the formulae:

21. (Original) The compound of claim 1, wherein R<sup>8</sup> and R<sup>9</sup>, taken together, form a group having one of the formulae:

- 22. (Original) The compound of claim 1, wherein r is 0 or 1.
- 23. (Original) The compound of claim 19, 20, or 21, wherein r is 1, 2, or 3, and each occurrence of halogen,  $C_{1-4}$ alkyl,  $-(R)_2$ , -OR, -SR,  $-SO_2N(R)_2$ ,  $-N(R)SO_2R$ , -N(R)COR,  $-N(R)_2$ ,  $-CH_2OR$ ,  $-CH_2N(R)_2$ , or  $-CH_2SR$ .

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24. (Original) The compound of claim 19, 20, or 21, wherein t is 0, Y is a bond, and R<sup>8</sup> is an optionally substituted heteroaryl moiety selected from one of groups b through r.

- 25. (Original) The compound of claim 24, wherein R<sup>8</sup> is an optionally substituted heteroaryl group b-i, k-i, or l-i.
- 26. (Original) The compound of claim 1, wherein t is 1, Y is O, S or NR<sup>9</sup>, and R<sup>8</sup> is optionally substituted phenyl.
- 27. (Original) The compound of claim 1, wherein t is 0 or 1, Y is NR<sup>9</sup>, and R<sup>8</sup> and R<sup>9</sup>, taken together form an optionally substituted group selected from s, u or v.
- 28. (Previously presented) The compound of claim 1, wherein Z<sup>3</sup> or Z<sup>5</sup> is CR<sup>3</sup> or CR<sup>5</sup>, respectively, and R<sup>3</sup> or R<sup>5</sup> is NRC(O)R<sup>7</sup>, wherein R<sup>7</sup> is (CH<sub>2</sub>)<sub>1</sub>-Y-R<sup>8</sup>, wherein t is 0, 1 or 2, wherein Y is a bond or is O, S, NR<sup>9</sup>, -OCH<sub>2</sub>-, -SCH<sub>2</sub>, -NR<sup>9</sup>CH<sub>2</sub>, O(CH<sub>2</sub>)<sub>2</sub>-, -S(CH<sub>2</sub>)<sub>2</sub>, or -NR<sup>9</sup>(CH<sub>2</sub>)<sub>2</sub>, and wherein R<sup>8</sup> is Ar<sup>2</sup>, or R<sup>8</sup> and R<sup>9</sup>, taken together with the nitrogen atom, form a 5-8 membered heterocyclyl or heteroaryl ring having 1-3 heteroatoms independently selected from nitrogen, oxygen or sulfur, and compounds have the formula II-A:

$$\begin{array}{c|c} R^{1} & \text{NH} \\ N & N \\ R^{Y}_{n}(U) & Z^{2} & N \\ R^{X}(T)_{m} & Z^{6} & B \\ Z^{5} & Z^{4} & O \end{array}$$

II-A

29. (Previously presented) The compound of claim 28, wherein ring B is selected from i, ii, iii, iv, v, vii, viii, ix, x, xi, xii, or xiii and compounds have one of formulas

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II-A-i, II-A-ii, II-A-iii, II-A-iv, II-A-v, II-A-viii, II-A-ix, II-A-x, II-A-xi, II-A-xii, or II-A-xiii:

$$\begin{array}{c|c}
R^1 & NH \\
N & N & R^2 \\
R^1 & NH \\
R^2 & R & R^4
\end{array}$$

II-A-i

II-A-iii

$$\begin{array}{c|cccc}
R^1 & NH & & & \\
N & N & R^2 & R & & \\
R^Y_n(U) & & & & & & \\
R^X(T)_m & & & & & & \\
R^5 & & & & & & \\
\end{array}$$

II-A-v

II-A-vii

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II-A-viii

II-A-ix

$$\begin{array}{c|c}
R^1 & NH \\
N & N & R^2 & R \\
R^Y_n(U) & B & N & Q
\end{array}$$

 $\Pi$ -A-x

II-A-xi

$$\begin{array}{c|c}
R^{1} & NH \\
N & N & R \\
R^{Y}_{n}(U) & N & R \\
R^{X}(T)_{m} & N & R \\
N & N & R^{4}
\end{array}$$

II-A-xii

II-A-xiii

## 30. (Canceled)

31. (Previously presented) The compound of claim 1, wherein  $Z^4$  is  $CR^4$ , and  $R^4$  is  $NRC(O)R^7$ , wherein  $R^7$  is  $(CH_2)_1-Y-R^3$ , wherein t is 0, 1 or 2, wherein Y is a bond or is O, S,  $NR^9$ ,  $-OCH_2$ -,  $-SCH_2$ ,  $-NR^9CH_2$ ,  $O(CH_2)_2$ -,  $-S(CH_2)_2$ , or  $-NR^9(CH_2)_2$ , and

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wherein R<sup>8</sup> is Ar<sup>2</sup>, or R<sup>8</sup> and R<sup>9</sup>, taken together with the nitrogen atom, form a 5-8 membered heterocyclyl or heteroaryl ring having 1-3 heteroatoms independently selected from nitrogen, oxygen or sulfur, and compounds have formula II-B:

$$R^{1} N +$$

$$N =$$

$$R^{Y}_{n}(U)$$

$$R^{X}(T)_{m}$$

$$Z^{2} Z^{3}$$

$$Z^{5}$$

$$R$$

$$R^{7}$$

II-B

32. (Previously presented) The compound of claim 31, wherein ring B is selected from i, ii, iii, iv, vi, viii, ix, xii, or xiv and compounds have one of formulas II-B-i, II-B-ii, II-B-iii, II-B-iv, II-B-vi, II-B-viii, II-B-xii, or II-B-xiv:

II-B-i

П-В-іі

II-B-iii

II-B-iv

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II-B-vi

II-B-viii

$$\begin{array}{c|c}
R^1 & NH \\
N & N & R^2 \\
R^{Y}_{n}(U) & B & R^3 \\
R^{X}(T)_{m} & R^6 & N & R^7
\end{array}$$

II-B-ix

II-B-xii

∏-B-xiv

## 33. (Canceled)

34. (Previously presented) The compound of claim 1, wherein  $Z^3$  or  $Z^5$  is  $CR^3$  or  $CR^5$ , respectively, and  $R^3$  or  $R^5$  is  $C(C^1)N(R)(R^7)$ , wherein  $R^7$  is  $(CH_2)_{r-}Y-R^8$ , wherein t is 0, 1 or 2, wherein Y is a bond or is O, S,  $NR^9$ , -OCH<sub>2</sub>-, -SCH<sub>2</sub>, -NR<sup>9</sup>CH<sub>2</sub>, O(CH<sub>2</sub>)<sub>2</sub>-, -S(CH<sub>2</sub>)<sub>2</sub>, or -NR<sup>9</sup>(CH<sub>2</sub>)<sub>2</sub>, and wherein  $R^8$  is  $Ar^2$ , or  $R^8$  and  $R^9$ , taken

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together with the nitrogen atom, form a 5-8 membered heterocyclyl or heteroaryl ring having 1-3 heteroatoms independently selected from nitrogen, oxygen or sulfur and compounds have formula **II-C**:

$$R^{1} NH$$

$$R^{Y}_{n}(U) \xrightarrow{R^{X}(T)_{r_{1}}} Z^{2} \xrightarrow{II-C} R^{X}$$

35. (Previously presented) The compound of claim 34, wherein ring B is selected from i, ii, iii, iv, v, vii, viii, ix, x, xi, xii, or xiii and compounds have one of formulas II-C-i, II-C-ii, II-C-iii, II-C-iv, II-C-v, II-C-vii, II-C-viii, II-C-x, II-C-xi, II-C-xii, or II-C-xiii:

$$\begin{array}{c|c}
R^1 & \text{NH} \\
N & N & R^2 & O \\
R^1 & N & R^2 & O \\
R^2 & N & R^3 & R^4 & R
\end{array}$$

II-C-ii

R<sup>1</sup>
NH
N O O N
R<sup>7</sup>
R<sup>7</sup>
R<sup>1</sup>
R<sup>8</sup>
R<sup>8</sup>
R<sup>7</sup>

II-C-iii

II-C-iv

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$$\begin{array}{c|cccc}
R^1 & & & & & \\
N & & & & & & \\
N & & & & & & \\
R^Y_n(U) & & & & & & \\
R^X(T)_m & & & & & & \\
R^5 & & & & & \\
\end{array}$$

II-C-v

$$\begin{array}{c|c}
R^{1} & NH \\
N & N & O \\
R^{Y}_{n}(U) & R^{X}(T)_{m} & R^{6} & R^{4}
\end{array}$$

II-C-viii

II-C-ix

$$\begin{array}{c|c}
R^1 & NH \\
N & N & R^2 & O \\
R^1 & NH & R^2 & O \\
R^2 & NH & R^2 & O \\
R^3 & R^4 & R^5 & R^5
\end{array}$$

$$\begin{array}{c|c}
R^1 & NH \\
N & N & R^2 & O \\
R^1 & NH & R^2 & O \\
R^2 & R^3 & R^4
\end{array}$$

II-C-x

П-С-хі

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II-C-xii

II-C-xiii

36. (Canceled)

37. (Previously presented) The compound of claim 1, wherein  $Z^4$  is  $CR^4$ , and  $R^4$  is  $C(O)N(R)(R^7)$ , wherein  $R^7$  is  $(CH_2)_{t^-}Y \cdot R^8$ , wherein t is 0, 1 or 2, wherein Y is a bond or is O, S,  $NR^9$ ,  $-OCH_2$ -,  $-SCH_2$ ,  $-NR^9CH_2$ ,  $O(CH_2)_2$ -,  $-S(CH_2)_2$ , or  $-NR^9(CH_2)_2$ , and wherein  $R^8$  is  $Ar^2$ , or  $R^8$  and  $R^9$ , taken together with the nitrogen atom, form a 5-8 membered heterocyclyl or heteroaryl ring having 1-3 heteroatoms independently selected from nitrogen, oxygen or sulfur and compounds have formula II-D:

$$R^{1} \cap H$$

$$R^{Y}_{n}(U) \cap Z^{2} \cap Z^{3} \cap R^{7}$$

$$H-D$$

38. (Previously presented) The compound of claim 37, wherein ring B is selected from i, ii, iii, iv, vi, viii, ix, xii, or xiv and compounds have one of formulas Π-D-i, Π-D-ii, Π-D-iii, Π-D-iv, Π-D-vi, Η-D-viii, Η-D-ix, Π-D-xii, or Π-D-xiv:

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$$R^1$$
 $NH$ 
 $R^2$ 
 $R^3$ 
 $R^3$ 
 $R^3$ 
 $R^4$ 
 $R^5$ 
 $R^5$ 

II-D-i

II-D-ii

$$\begin{array}{c|c} R^1 & NH \\ N & N & R^2 \\ R^Y_{n}(U) & & B & R^3 \\ R^X(T)_m & & N & N & N \end{array}$$

Π-D-iii

II-D-iv

II-D-vi

∏-D-viii

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П-D-ix

II-D-xii

II-D-xiv

## 39. (Canceled)

40. (Previously presented) The compound of claim 1, where R<sup>1</sup> is optionally substituted phenyl and ring B is an optionally substituted phenyl group and compounds have the general formula IV:

$$q(R^{Z}Z)$$

NH

NA

N

R

R

R

R

R

R

R

R

IV

41. (Previously presented) The compound of claim 40, wherein, R<sup>3</sup> is NRCOR<sup>7</sup> and compounds have the general formula IV-A-(i):

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$$q(R^{Z}Z)$$

NH

 $R^{Y}_{n}(U)$ 
 $R^{X}(T)_{m}$ 
 $R^{B}$ 
 $R^{A}$ 
 $R^{A}$ 

42. (Previously presented) The compound of claim 40, wherein R<sup>4</sup> is NRCOR<sup>7</sup> and compounds have the general formula IV-B-(i):

43. (Previously presented) The compound of claim 40, wherein R<sup>3</sup> is CONRR<sup>7</sup> and compounds have the general formula IV-C-(i):

$$q(R^{Z}Z)$$
 $NH$ 
 $R^{Y}_{n}(U)$ 
 $R^{X}(T)_{m}$ 
 $R^{\theta}$ 
 $R^{\theta}$ 
 $R^{\phi}$ 
 $R^{\phi}$ 

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44. (Previously presented) The compound of claim 40, wherein R<sup>4</sup> is CONRR<sup>7</sup> and compounds have the general formula IV-D-(i):

$$\begin{array}{c|c} & & & & \\ & & & & \\ & & & & \\ R^{Y}_{n}(U) & & & & \\ & & & & \\ R^{X}(T)_{m} & & & \\ & & & \\ R^{6} & & & \\ & & & \\ R^{5} & & \\ \end{array} \begin{array}{c} & & \\ R^{3} \\ & & \\ R^{7} \end{array}$$

IV-D-(i)

45. (Currently amended) The compound of claim 40, wherein R<sup>1</sup> is optionally substituted phenyl, ring A is pyrimidinyl, ring B is phenyl, and R<sup>2</sup>, R<sup>5</sup>, and R<sup>6</sup> are each hydrogen, and compounds have the general <u>formula</u>VI:

- (Currently amended) The compound of claim 40 or 45, wherein
  (a) q is 0 or 1 and ZR<sup>Z</sup> is -NH<sub>2</sub>, -OH, C<sub>1-4</sub>alkoxy, or -SO<sub>2</sub>NH<sub>2</sub>;
  (b) R<sup>3</sup> is NRCOR<sup>7</sup>, wherein R<sup>7</sup> is (CH<sub>2</sub>)<sub>1</sub>-Y-R<sup>8</sup>, and t is 0, Y is a bond, and R<sup>8</sup> is phenyl (a), or is an optionally substituted heteroaryl moiety selected from one of groups b through r, and wherein r is 0 or 1, and WR<sup>W</sup> substituents are halogen. C<sub>1-4</sub>alkyl, -(R)<sub>2</sub>, -OR, -SR, -SO<sub>2</sub>N(R)<sub>2</sub>, -N(R)SO<sub>2</sub>R, -N(R)COR, -N(R)<sub>2</sub>, -CH<sub>2</sub>OR, -CH<sub>2</sub>N(R)<sub>2</sub>, or -CH<sub>2</sub>SR; and
  (c) R<sup>4</sup> is hydrogen.
- 47. (Previously presented) The compound of claim 40 or 45, wherein:

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- (a) q is 0 or 1 and  $\mathbb{ZR}^2$  is -NH<sub>2</sub>, -OH,  $\mathbb{C}_{1-4}$  alkoxy, or -SO<sub>2</sub>NH<sub>2</sub>;
- (b) R<sup>3</sup> is CONRR<sup>7</sup>, wherein R<sup>7</sup> is (CH<sub>2</sub>)<sub>t</sub>-Y-R<sup>8</sup>, and t is 0, Y is a bond, and R<sup>8</sup> is phenyl (a) or is an optionally substituted heteroaryl moiety selected from one of groups b through r, and wherein r is 0 or 1, and WR<sup>W</sup> substituents are halogen, C<sub>1.4</sub>alkyl, -(R)<sub>2</sub>, -OR, -SR, -SO<sub>2</sub>N(R)<sub>2</sub>, -N(R)SO<sub>2</sub>R, -N(R)COR, -N(R)<sub>2</sub>, -CH<sub>2</sub>OR, -CH<sub>2</sub>N(R)<sub>2</sub>, or -CH<sub>2</sub>SR, and (c) R<sup>4</sup> is hydrogen.
- .
- 48. (Previously presented) The compound of claim 40 or 45, wherein:
  - (a) q is 0 or 1 and  $\mathbb{ZR}^{\mathbb{Z}}$  is -NH<sub>2</sub>, -OH,  $\mathbb{C}_{1}$ -alkoxy, or -S(O)<sub>2</sub>NH<sub>2</sub>;
  - (b)  $R^4$  is  $NRCOR^7$ , wherein  $R^7$  is  $(CH_2)_t$ -Y- $R^8$ , and t is 0, Y is a bond, and  $R^8$  is phenyl (a) or an optionally substituted heteroaryl moiety selected from one of groups **b** through **z**, and wherein **r** is 0 or 1, and  $WR^W$  substituents are halogen,  $C_{1-4}$ alkyl,  $-(R)_2$ , -OR, -SR,  $-SO_2N(R)_2$ ,  $-N(R)SO_2R$ , -N(R)COR,  $-N(R)_2$ ,  $-CH_2OR$ ,  $-CH_2N(R)_2$ , or  $-CH_2SR$ ; and
  - (c) R<sup>3</sup> is hydrogen.
- 49. (Previously presented) The compound of claim 40 or 45, wherein:
  - (a) q is 0 or 1 and ZR<sup>Z</sup> is -NH<sub>2</sub>, -OH, C<sub>1-4</sub>alkoxy, or -S(O)<sub>2</sub>NH<sub>2</sub>;
  - (b)  $R^4$  is  $CONRR^7$ , wherein  $R^7$  is  $(CH_2)_t$ -Y- $R^8$ , and t is 0, Y is a bond, and  $R^8$  is phenyl (a) or an optionally substituted heteroaryl moiety selected from one of groups b through z, and wherein r is 0 or 1, and  $WR^W$  substituents are halogen,  $C_{1-4}$ alkyl,  $-(R)_2$ , -OR, -SR,  $-SO_2N(R)_2$ ,  $-N(R)SO_2R$ , -N(R)COR,  $-N(R)_2$ ,  $-CH_2OR$ ,  $-CH_2N(R)_2$ , or  $-CH_2SR$ ; and
  - (c) R<sup>3</sup> is hydrogen.
- 50. (Previously presented) The compound of claim 40 or 45, wherein:
  - (a) q is 0 or 1 and  $\mathbb{Z}\mathbb{R}^Z$  is -NH<sub>2</sub>, -OH, C<sub>1-4</sub>alkoxy, or -S(O)<sub>2</sub>NH<sub>2</sub>;

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(b) R<sup>3</sup> is NRCOR<sup>7</sup>, wherein R<sup>7</sup> is (CH<sub>2</sub>)<sub>t</sub>-Y-R<sup>8</sup>, and t is 0 or 1, Y is NR<sup>9</sup>, and R<sup>8</sup> and R<sup>9</sup>, taken together with the nitrogen atom, form a group selected from s, t, u, or v, and wherein r is 0 or 1, and WR<sup>W</sup> substituents are halogen, C<sub>1-4</sub>alkyl, -(R)<sub>2</sub>, -OR, -SR, -SO<sub>2</sub>N(R)<sub>2</sub>, -N(R)SO<sub>2</sub>R, -N(R)COR, -N(R)<sub>2</sub>, -CH<sub>2</sub>OR, -CH<sub>2</sub>N(R)<sub>2</sub>, or -CH<sub>2</sub>SR; and

- (c) R<sup>4</sup> is hydrogen.
- 51. (Previously presented) The compound of claim 40 or 45, wherein:
  - (a) q is 0 or 1 and  $\mathbb{ZR}^2$  is -NH<sub>2</sub>, -OH,  $\mathbb{C}_{1-4}$  alkoxy, or -S(O)<sub>2</sub>NH<sub>2</sub>;
  - (b)  $R^3$  is  $CONRR^7$ , wherein  $R^7$  is  $(CH_2)_t$ -Y- $R^8$ , and t is 0 or 1, Y is  $NR^9$ , and  $R^8$  and  $R^9$ , taken together with the nitrogen atom, form a group selected from s, t, u, or v, and wherein r is 0 or 1, and  $WR^W$  substituents are halogen,  $C_{1-4}$ alkyl, -(R)<sub>2</sub>, -OR, -SR, -SO<sub>2</sub>N(R)<sub>2</sub>, -N(R)SO<sub>2</sub>R, -N(R)COR, -N(R)<sub>2</sub>, -CH<sub>2</sub>OR, -CH<sub>2</sub>N(R)<sub>2</sub>, or -CH<sub>2</sub>SR; and
  - (c) R<sup>4</sup> is hydrogen.
- 52. (Previously presented) The compound of claim 40 or 45, wherein:
  - (a) q is 0 or 1 and  $\mathbb{ZR}^{\mathbb{Z}}$  is -NH<sub>2</sub>, -OH, C<sub>1.4</sub>alkoxy, or -S(O)<sub>2</sub>NH<sub>2</sub>;
  - (b) R<sup>4</sup> is NRCOR<sup>7</sup>, wherein R<sup>7</sup> is (CH<sub>2</sub>)<sub>t</sub>-Y-R<sup>8</sup>, and t is 0 or 1, Y is NR<sup>9</sup>, and R<sup>8</sup> and R<sup>9</sup>, taken together with the nitrogen atom, form a group selected from s, t, u, or v, and wherein r is 0 or 1, and WR<sup>W</sup> substituents include halogen, C<sub>1-4</sub>alkyl, NH<sub>2</sub>, OH, SH, SO<sub>2</sub>NH<sub>2</sub>, C<sub>1-4</sub>alkoxy, C<sub>1</sub>. 4thioalkyl, CH<sub>2</sub>OR, CH<sub>2</sub>N(R)<sub>2</sub>, or CH<sub>2</sub>SR; and
  - (c) R<sup>3</sup> is hydrogen.
- 53. (Previously presented) The compound of claim 40 or 45, wherein:
  - (a) q is 0 or 1 and  $\mathbb{ZR}^{\mathbb{Z}}$  is -NH<sub>2</sub>, -OH, C<sub>1-4</sub>alkoxy, or -S(O)<sub>2</sub>NH<sub>2</sub>;
  - (b) R<sup>4</sup> is CONRR<sup>7</sup>, wherein R<sup>7</sup> is (CH<sub>2</sub>)<sub>t</sub>-Y-R<sup>8</sup>, and t is 0 or 1, Y is NR<sup>9</sup>, and R<sup>8</sup> and R<sup>9</sup>, taken together with the nitrogen atom, form a group

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selected from s, t, u, or v, and wherein r is 0 or 1, and WR<sup>W</sup> substituents are halogen,  $C_{1-4}$ alkyl,  $(R)_2$ , -OR, -SR,  $-SO_2N(R)_2$ ,  $-N(R)SO_2R$ , -N(R)COR,  $-N(R)_2$ ,  $-CH_2OR$ ,  $-CH_2N(R)_2$ , or  $-CH_2SR$ ; and (c)  $R^3$  is hydrogen.

54. (Previously presented) The compound of claim 1, having one of the following structures:

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- 55. (Original) A pharmaceutical composition comprising a compound according to claim 1, and a pharmaceutically acceptable carrier, adjuvant, or vehicle.
- 56. (Currently amended) The composition of claim 55, further comprising an additional therapeutic agent selected from a chemotherapeutic or anti-proliferative agent, a treatment for Alzheimer's Discase, a treatment for Parkinson's Discase, an agent for treating Multiple Sclerosis (MS), a treatment for asthma, an agent for treating schizophrenia, an anti-inflammatory agent [[,]] or an immunomodulatory or immunosuppressive agent, a neurotrophic factor, an agent for treating cardiovascular disease, an agent for treating destructive bone disorders, an agent for treating liver disease, an agent for treating a blood disorder, or an agent for treating an immunodeficiency disorder.
- 57. (Currently amended) A method of inhibiting JAK kinase activity in a biological sample <u>in vitro</u> or a patient, comprising the step of contacting said biological sample or patient with:
  - a) the composition of claim 55; or
  - b) the compound of claim 1.
- 58. (Currently amended) A method of treating or lessening the severity of a disease or disorder selected from <u>rheumatoid arthritis</u>, <u>allergic or type I</u> hypersensitivity reaction, asthma, familial amyotrophic lateral sclerosis (FALS) or

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transplant rejection, an immune response, an autoimmune disease, a neurodegenerative disorder, or a solid or hematologic malignancy comprising administering to a patient in need thereof a compound of claim 1 or a composition of claim 55.

59. (Canceled)